Serial No. 09/581,772 Docket No. PP01388.202

a polymer selected from the group consisting of a poly( $\alpha$  -hydroxy acid), a polyhydroxy butyric acid, a polycaprolactone, a polyorthoester, a polyanhydride, and a polycyanoacrylate; and

a cationic or anionic detergent.

- 2. (Amended) The microparticle of claim 1, further comprising a first biologically active macromolecule adsorbed on the surface thereof, wherein the first biologically active macromolecule is at least one member selected from the group consisting of a polypeptide, a polynucleotide, a polynucleoside, an antigen, a pharmaceutical, a hormone, an enzyme, a transcription or translation mediator, an intermediate in a metabolic pathway, an immunomodulator, and an adjuvant.
- 3. (Amended) The microparticle of claim 2, further comprising a second biologically active macromolecule encapsulated within said microparticle, wherein the second biologically active macromolecule is at least one member selected from the group consisting of a polypeptide, a polynucleotide, a polynucleoside, an antigen, a pharmaceutical, a hormone, an enzyme, a transcription or translation mediator, an intermediate in a metabolic pathway, an immunomodulator, and an adjuvant.



- 9. (Amended) The microparticle of any of claims 2-7, wherein the first biologically active macromolecule is an antigen selected from the group consisting of gp120, gp160, p24gag, p55gag, and Influenza A hemagglutinin antigen.
- 10. (Amended) The microparticle of any of claims2-7, wherein the first biologically active macromolecule is a polynucleotide which encodes gp120.
- 11. (Amended) The microparticle of any of claims 3-7, 9 and 10, wherein the second biologically active macromolecule is an adjuvant.
- 12. (Amended) The microparticle of claim 11, wherein the adjuvant is an aluminum salt.

13. (Amended) A microparticle composition comprising a microparticle of any of claims 1-7 and 9-12 and a pharmaceutically acceptable excipient.



- 14. (Amended) A microparticle composition comprising a microparticle according to any of claims 1-7, 9, 10 and 13, further comprising an adjuvant.
- 17. (Amended) A method of producing a microparticle having an adsorbent surface, said method comprising the steps of:
- (a) dispersing a mixture of a polymer solution and a cationic or anionic detergent, wherein the polymer solution comprises a polymer selected from the group consisting of a poly(α -hydroxy acid), a polyhydroxy butyric acid, a polycaprolactone, a polyorthoester, a polyanhydride, and a polycyanoacrylate, wherein the polymer is present at a concentration of about 1% to about 30% in an organic solvent, and wherein the detergent is present in the mixture at a weight to weight detergent to polymer ratio of from about 0.00001:1 to about 0.1:1; and
  - (b) removing the organid solvent from the emulsion.
- 21. (Amended) The method of any of claims 17-19 wherein the detergent is present at a weight to weight detergent to polymer ratio of from about 0.0001:1 to about 0.01:1.
- 22. (Amended) The method of any of claims 17-19 wherein the detergent is present at a weight to weight detergent to polymer ratio of from about 0.001:1 to about 0.01:1.
- 23. (Amended) The method of any of claims 17-19 wherein the detergent is present at a weight to weight detergent to polymer ratio of from about 0.005:1 to about 0.01:1.
- 24. (Amended) The method of any of claims 17-19 and 21-23, wherein the microparticle comprises a poly(α hydroxy acid) selected from the group consisting of poly(L-lactide), poly(D,L-lactide) and poly(D,L-lactide-co-glycolide).



- 27. (Amended) A method of producing a microparticle having an adsorbent surface to which a biologically active macromolecule has been adsorbed, said method comprising the steps of:
- (a) emulsifying a mixture of a polymer solution and a cationic or anionic detergent to form an emulsion, wherein the polymer solution comprises a polymer selected from the group consisting of a poly( $\alpha$  -hydroxy acid), a polyhydroxy butyric acid, a polycaprolactone, a polyorthoester, a polyanhydride, and a polycyanoacrylate, wherein the polymer is present at a concentration of about 1% to about 30% in an organic solvent, and wherein the detergent is present in the mixture at a weight to weight detergent to polymer ratio of from about 2.00001:1 to about 0.1:1;
- (b) removing the organic solvent from the emulsion, to form said microparticle having the adsorbent surface; and
  - (c) adsorbing the macromolecule to the surface of the microparticle.

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29. (Amended) The method of any of claims 27-28, wherein the macromolecule is an antigen selected from the group consisting of gp120, gp160, p24gag, p55gag and Influenza A hemagglutinin antigen.

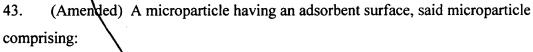
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34. (Amended) A miore particle made according to the method of any of claims 17-19 and 21-33.

- 36. (Amended) A method of producing a microparticle composition comprising a microparticle having an adsorbent surface to which a biologically active macromolecule has been adsorbed, said method comprising the steps of:
- (a) emulsifying a mixture of a polymer solution and a cationic or anionic detergent to form an emulsion, wherein the polymer solution comprises a polymer selected from the group consisting of a poly( $\alpha$  -hydroxy acid), a polyhydroxy butyric acid, a polycaprolactone, a polyorthoester, a polyanhydride, and a polycyanoacrylate, wherein the polymer is present at a concentration of about 1% to about 30% in an organic solvent, and wherein the detergent is present at a weight to weight detergent to polymer ratio of from about 0.00001:1 to about 0.1:1;

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- (b) removing the organic solvent from the emulsion, to form said microparticle having the adsorbent surface;
  - (c) adsorbing the macromolecule to the surface of the microparticle; and
- (d) combining the microparticle having the adsorbed macromolecule from step (c) with a pharmaceutically acceptable excipient to form said microparticle composition.



a biodegradable polymer; and

a cationic or artionic detergent.

44. (Amended) The microparticle of claim 43, further comprising a first biologically active macromolecule adsorbed on the surface thereof, wherein the first biologically active macromolecule is at least one member selected from the group consisting of a polypeptide, a polynucleotide, a polynucleoside, an antigen, a pharmaceutical, a hormone, an enzyme, a transcription or translation mediator, an intermediate in a metabolic pathway, an immunomodulator, and an adjuvant.

## Please add new claims 52-68 as follows:

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- 52. (Newly Added) The microparticle of claim 7, wherein the first biologically active macromolecule is a polypeptide.
- 53. (Newly Added) The microparticle of claim 52, wherein the first biologically active macromolecule is a polypeptide antigen selected from the group consisting of HIV antigens, hepatitis C virus antigens, and influenza A virus antigens.
- 54. (Newly Added) The microparticle of claim 6, wherein the first biologically active macromolecule is a polynucleotide.

55. (Newly Added) The microparticle of claim 54, wherein the polynucleotide encodes an antigen.

- 56. (Newly Added) The microparticle of claim 55, wherein the polynucleotide encoding the antigen is a plasmid DNA molecule.
- 57. (Newly Added) The microparticle of claim 55, wherein the antigen is selected from the group consisting of HIV antigens, hepatitis C virus antigens, and influenza A virus antigens.
- 58. (Newly Added) The microparticle of claim 6, wherein the cationic detergent is hexadecyltrimethylammonium bromide.
- 59. (Newly Added) The microparticle of claim 7, wherein the anionic detergent is sodium dodecyl sulfate.
- 60. (Newly Added) The method of claim 27, wherein the detergent is an anionic detergent.
- 61. (Newly Added) The method of claim 60, wherein the macromolecule is a polypeptide.
- 62. (Newly Added) The method of claim 61, wherein the polypeptide is a polypeptide antigen selected from the group consisting of HIV antigens, hepatitis C virus antigens, and influenza A virus antigens.
- 63. (Newly Added) The method of claim 27, wherein the detergent is a cationic detergent.
- 64. (Newly Added) The method of claim 63, wherein the macromolecule is a polynucleotide.

- 65. (Newly Added) The method of claim 64, wherein the polynucleotide encodes an antigen.
- 66. (Newly Added) The method of claim 65, wherein the polynucleotide encoding the antigen is plasmid DNA.
- 67. (Newly Added) The method of claim 65, wherein the antigen is selected from the group consisting of HIV antigens, hepatitis C virus antigens, and influenza A virus antigens.
- 68. (Newly Added) Use of a microparticle composition of claim 51, wherein said immune response comprises a CNL immune response.

Please cancel claims 8 and 20 without prejudice or disclaimer.

## **STATUS OF CLAIMS:**

Claims 1-7, 9-19 and 21-68 are pending herein.

Claims 52-68 have been added, and claims 8 and 20 have been cancelled without prejudice or disclaimer herein.

## **REMARKS**

## A. Claim Amendments

In view of the above amendment, claims 1-7, 9-19 and 21-68 are presently pending.

A separate sheet entitled "Version with Markings to Show Changes Made" is provided to illustrate the amendments made to claims 1-3, 9-14, 17, 21-24, 27, 29, 34, 36, 43 and 44, the cancellation of claims 8 and 20, and the addition of claims 52-68.